## AMENDMENT TO THE CLAIMS:

Please cancel claims 466 and 482-486 without prejudice or disclaimer and amend the claims as shown below.

## 1-432. (Previously cancelled)

- 433. (Currently amended) A composition comprising at least two types of nanoparticles having oligonucleotides attached thereto, at least a portion of the oligonucleotides on a first type of nanoparticles having a sequence complementary to a sequence of a first portion of a target nucleic acid, and at least a portion of the oligonucleotides on a second type of nanoparticles having a sequence complementary to a sequence of a second portion of the nucleic acid, wherein at least some the oligonucleotides bound to the nanoparticles comprise at least one type of recognition oligonucleotides, each type of recognition oligonucleotides comprising a sequence complementary to at least one portion of a sequence of the target nucleic acid; and a type of diluent oligonucleotides.
- 434. (Previously added) The composition of Claim 433 wherein the target nucleic acid is viral RNA or DNA.
- 435. (Previously added) The composition of Claim 433 wherein the target nucleic acid is a gene associated with a disease.
- 436. (Previously added) The composition of Claim 433 wherein the target nucleic acid is a bacterial DNA.
- 437. (Previously added) The composition of Claim 433 wherein the target nucleic acid is a fungal DNA.

- 438. (Previously added) The composition of Claim 433 wherein the target nucleic acid is a synthetic DNA, a synthetic RNA, a structurally-modified natural or synthetic RNA, or a structurally-modified natural or synthetic DNA.
- 439. (Previously added) The composition of Claim 433 wherein the target nucleic acid is from a biological source.
- 440. (Previously added) The composition of Claim 433 wherein the target nucleic acid is a product of a polymerase chain reaction amplification.
- 441. (Previously added) The composition of Claim 433 wherein the target nucleic acid comprises one or more types of linking oligonucleotides.
- 442. (Previously added) The composition of Claim 441 wherein the linking oligonucleotide is single-stranded.
- 443. (Previously added) The composition of Claim 441 wherein the linking oligonucleotide is double-stranded and has overhanging ends.
- 444. (Previously added) The composition of Claim 441 wherein the linking oligonucleotide is a triple-stranded oligonucleotide connector.
- 445. (Previously added) The composition of Claim 441 wherein the linking oligonucleotide comprises a peptide nucleic acid chain.
- 446. (Previously added) The composition of Claim 433 wherein the oligonucleotides are attached to the nanoparticles in a stepwise ageing process comprising (i) contacting the oligonucleotides with the nanoparticles in a first aqueous solution for a period of time sufficient to allow some of the oligonucleotides to bind to the nanoparticles; (ii) adding at least one salt to the first aqueous solution to create a second aqueous solution; and (iii) contacting the

oligonucleotides and nanoparticles in the second aqueous solution for an additional period of time to enable additional oligonucleotides to bind to the nanoparticles.

- 447. (Previously added) The composition of Claim 448 wherein the second aqueous solution has an ionic strength sufficient to overcome at least partially the electrostatic attraction or repulsion of the oligonucleotides for the nanoparticles and the electrostatic repulsion of the oligonucleotides to each other.
- 448. (Previously added) The composition of Claim 446 wherein the oligonucleotides and nanoparticles are contacted in aqueous solution for about 12 to about 24 hours.
- 449. (Previously added) The composition of Claim 446 wherein salt is added to the aqueous solution to form the aqueous salt solution which is buffered at pH 7.0 and which contains about 0.1 M NaCl.
- 450. (Previously added) The composition of Claim 446 wherein the oligonucleotides and nanoparticles are contacted in the aqueous salt solution for an additional 40 hours to increase the density of oligonucleotides bound to the nanoparticles.
- 451. (Previously added) The composition of Claim 449 wherein the salt is added to the first aqueous solution in a single addition.
- 452. (Previously added) The composition of Claim 449 wherein the salt is added gradually to the first aqueous solution over time.
- 453. (Previously added) The composition of Claim 449 wherein the salt is selected from the group consisting of consisting of sodium chloride, magnesium chloride, potassium chloride, ammonium chloride, sodium acetate, ammonium acetate, a combination of two or more of these salts, one of these salts in a phosphate buffer, and a combination of two or more these salts in a phosphate buffer.

- 454. (Previously added) The composition of Claim 453 wherein the salt is sodium chloride in a phosphate buffer.
- 455. (Previously added) The composition of Claim 433 wherein the oligonucleotides are present on a surface of the nanoparticles at a surface density of at least 10 picomoles/cm<sup>2</sup>.
- 456. (Previously added) The composition of Claim 455 wherein the oligonucleotides are present on the surface of the nanoparticles at a surface density of at least 15 picomoles/cm<sup>2</sup>.
- 457. (Previously added) The composition of Claim 456 wherein the oligonucleotides are present on the surface of the nanoparticles at a surface density from about 15 picomoles/cm<sup>2</sup> to about 40 picomoles/cm<sup>2</sup>.
- 458. (Previously added) The composition of Claim 433 wherein the nanoparticles are metal nanoparticles or semiconductor nanoparticles.
- 459. (Previously added) The composition of Claim 458 wherein the nanoparticles are gold nanoparticles.
- 460. (Previously added) The composition of Claim 458 wherein the semiconductor nanoparticles are made of CdSe/ZnS (core/shell).
- 461. (Previously added) The composition of Claim 433 wherein at least some of the oligonucleotides on the nanoparticles comprise at least one type of recognition oligonucleotides, each type of recognition oligonucleotides comprising a spacer portion and a recognition portion, the spacer portion being designed so that it is bound to the nanoparticles, the recognition portion having a sequence complementary to at least a portion of a sequence of the target nucleic acid.

- 462. (Previously added) The composition of Claim 461 wherein the spacer portion has a moiety covalently bound to it, the moiety comprising a functional group through which the spacer portion is bound to the nanoparticles.
- 463. (Previously added) The composition of Claim 461 wherein the spacer portion comprises at least about 10 nucleotides.
- 464. (Previously added) The composition of Claim 463 wherein the spacer portion comprises from about 10 to about 30 nucleotides.
- 465. (Previously added) The composition of Claim 461 wherein the bases of the nucleotides of the spacer portion are all adenines, all thymines, all cytosines, all uracils or all guanines.
- 466. (Currently cancelled) The composition of Claim 433 wherein at least some the oligonucleotides bound to the nanoparticles comprise at least one type of recognition oligonucleotides, each type of recognition oligonucleotides comprising a sequence complementary to at least one portion of a sequence of the target nucleic acid; and a type of diluent oligonucleotides.
- 467. (Currently amended) The composition of Claim [466] <u>433</u> wherein, each type of recognition oligonucleotides comprises a spacer portion and a recognition portion, the spacer portion being designed so that it is bound to the nanoparticles, the recognition portion having a sequence complementary to at least one portion of a sequence of the target nucleic acid.
- 468. (Previously added) The composition of Claim 467 wherein the spacer portion has a moiety covalently bound to it, the moiety comprising a functional group through which the spacer portion is bound to the nanoparticles.

- 469. (Previously added) The composition of Claim 467 wherein the spacer portion comprises at least about 10 nucleotides.
- 470. (Previously added) The composition of Claim 469 wherein the spacer portion comprises from about 10 to about 30 nucleotides.
- 471. (Previously added) The composition of Claim 467 wherein the bases of the nucleotides of the spacer portion are all adenines, all thymines, all cytosines, all uracils or all guanines.
- 472. (Previously added) The composition of Claim 467 wherein the diluent oligonucleotides contain about the same number of nucleotides as are contained in the spacer portions of the recognition oligonucleotides.
- 473. (Previously added) The composition of Claim 472 wherein the sequence of the diluent oligonucleotides is the same as that of the spacer portions of the recognition oligonucleotides.
- 474. (Currently amended) The composition of any one of Claims 433[,] or 461, [or 466] wherein the oligonucleotides are bound to the nanoparticles through sulfur linkages.
- 475. (Previously added) The composition of Claim 433 wherein in the presence of target nucleic acid and under hybridization conditions, he composition forms a complex with the target nucleic acid, the resulting complex having a sharp melting profile and increased melting temperature relative to a comparable complex without nanoparticles to allow for selective discrimination of any nucleotide insertion, deletion or mismatch in the target nucleic acid.
- 476. (Currently amended) A kit comprising the composition of claim 433 [having at least two types of nanoparticles having oligonucleotides attached thereto, at least a portion of the oligonucleotides on the first type of nanoparticles having a sequence complementary to the

sequence of a first portion of a nucleic acid, and at least a portion of the oligonucleotides on the second type of nanoparticles having a sequence complementary to the sequence of a second portion of the nucleic acid].

- 477. (Currently amended) The kit of Claim [433] <u>476</u> wherein the target nucleic acid comprises one or more types of linking oligonucleotides.
- 478. (Previously added) The kit of Claim 477 wherein the linking oligonucleotide is single-stranded.
- 479. (Previously added) The kit of Claim 477 wherein the linking oligonucleotide is double-stranded and has overhanging ends.
- 480. (Previously added) The kit of Claim 477 wherein the linking oligonucleotide is a triple-stranded oligonucleotide connector.
- 481. (Previously added) The kit of Claim 477 wherein the linking oligonucleotide comprises a peptide nucleic acid chain.
- 482. (Currently cancelled) A kit comprising a composition having at least two types of nanoparticles having oligonucleotides attached thereto, at least a portion of the oligonucleotides on the first type of nanoparticles having a sequence complementary to the sequence of a first portion of a linking oligonucleotide, and at least a portion of the oligonucleotides on the second type of nanoparticles having a sequence complementary to the sequence of a second portion of the nucleotide; and one or more types of linking oligonucleotides.
- 483. (Currently cancelled) The kit of Claim 477 wherein the linking oligonucleotide is single-stranded.

- 484. (Currently cancelled) The kit of Claim 477 wherein the linking oligonucleotide is double-stranded and has overhanging ends.
- 485. (Currently cancelled) The kit of Claim 477 wherein the linking oligonucleotide is a triple-stranded oligonucleotide connector.
- 486. (Currently cancelled) The kit of Claim 477 wherein the linking oligonucleotide comprises a peptide nucleic acid chain.